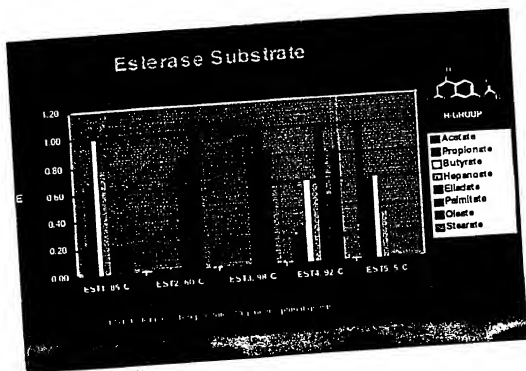
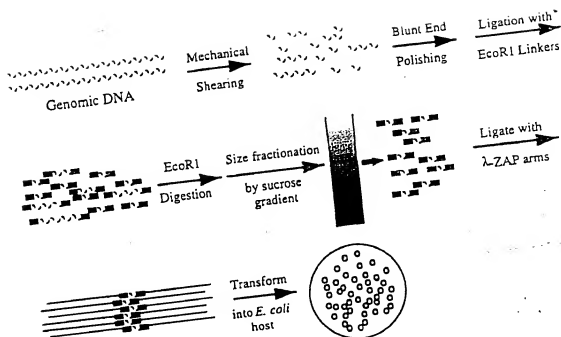


Figure 1



09636778.081100

Figure 2.



09635778.081107

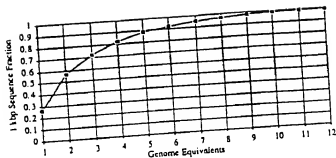


Figure 3.

09636779-081100

Cell sorting to screen for recombinant enzymes

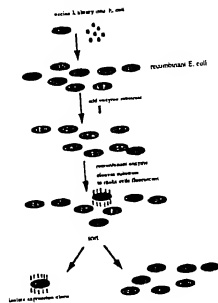
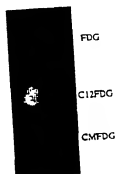


Figure 4.

0066778-081100

β -Gal clone with different substrates

- cells were stained with FDG, CMFDG or C12FDG, incubated for 30 min. at 70°C, spotted onto a slide and exposed to UV light.
- bright spot indicates staining of cells



E. coli expressing β -Gal from Sulfulobus spec. was grown over night. Cells were centrifuged and substrate was loaded with deionised water. After 5 min. cells were centrifuged and transferred into HEPES buffer and heated to 70°C for 30 min.. Cells were spotted onto a slide and exposed to UV light.

Figure 5

UOTTSO-8/29E960

Figure 6

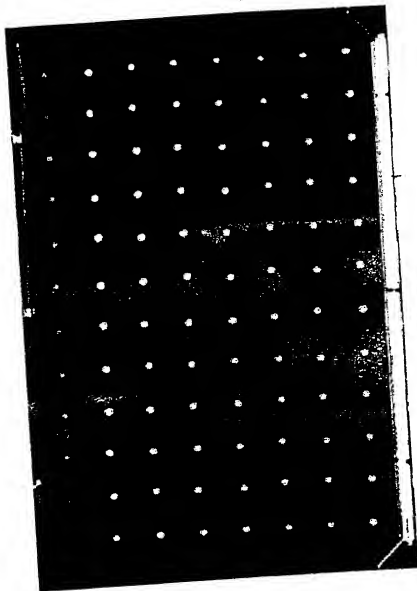
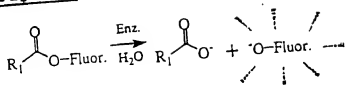


Figure 7



Principle type of fluorescence enzyme assay of deacylation.

0955779-08100

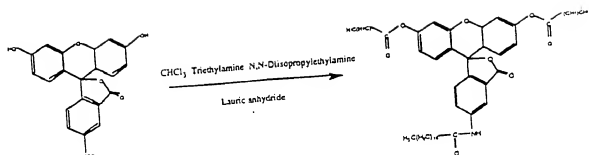
Figure 8



Staining of β -galactosidase clones from the hyperthermophilic archaeon *Sulfolobus solfataricus* expressed in *E.coli* using C_{12} -FDG as enzyme substrate.

0963678-00100

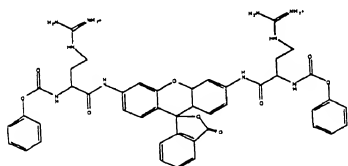
Figure 9



Synthesis of 5-dodecanoyl-aminofluorescein-di-dodecanoic acid

09635779-081100

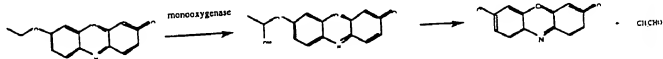
Figure 10



Rhodamine protease substrate.

00180*8729560

Figure 11



Compound and process that can be used in the detection of monooxygenases

0056678-08100

Combinatorial Enzyme Development

(Natural + Non-natural Evolution)

Desired
Enzyme

=

Improve
Nature

+

Search
Nature

Directed
Evolution

Select
Enzyme

Enzyme
Library

Enzyme 1

Enzyme 2

Enzyme 3

Enzyme 4

Enzyme 5

Enzyme 6

Enzyme 7

Enzyme 8

Enzyme 9

Enzyme 10

Enzyme n

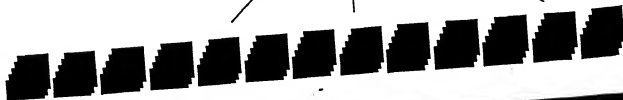
ID via High
Throughput
Screening

ID via Enzyme
Characterization

ID via Mutation /
Selection

New Enzyme

NA Library



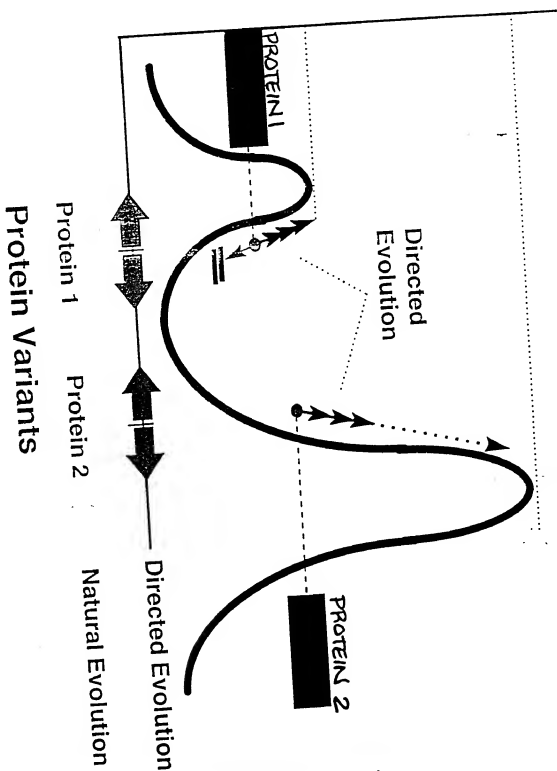
Bypassing Barriers to Directed Protein Evolution

(Barrier = Capacity limit of directed evolution system)

FIGURE 13

- STABILITY
- Solvent Stability
 - Expression Level
 - Buffer Compatibility
 - Process Compatibility

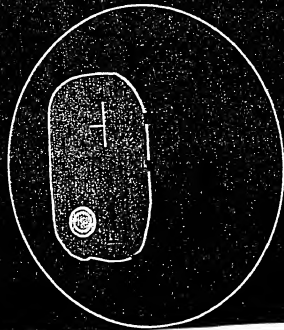
RELATIVE
ENZYME
ACTIVITY



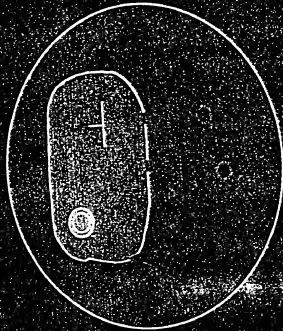
09536778-081100



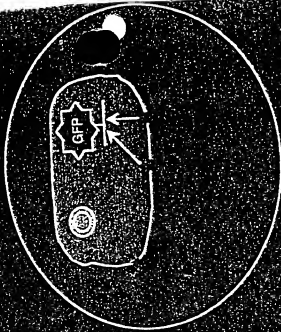
Co-Encapsulation Assay for Novel Bioactive Discovery



Co-encapsulation
Library + Eukaryote



Growth and expression
of small molecule from host



Receptor binding of small
molecule & GFP reporter

E = Eukaryotic assay organism L = Large insert library SM = Small molecule
GFP = Green Fluorescent Protein R = Eukaryotic receptor



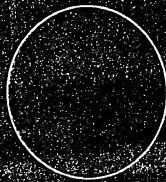
FACS Screening for Encapsulation

Test organisms

Pathway clones

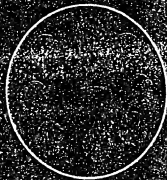


encapsulate

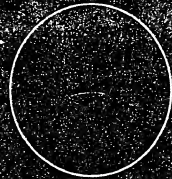


live/dead or other
activity stain

sort



bioactive expression
(e.g. live/dead, growth rate,
metabolic stains etc.)

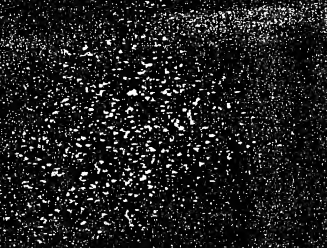




Development of Unicellular *Streptomyces* Strains for Ultra-High Throughput Screening



Streptomyces "diversa"
unicells



Streptomyces lividans
mycelia

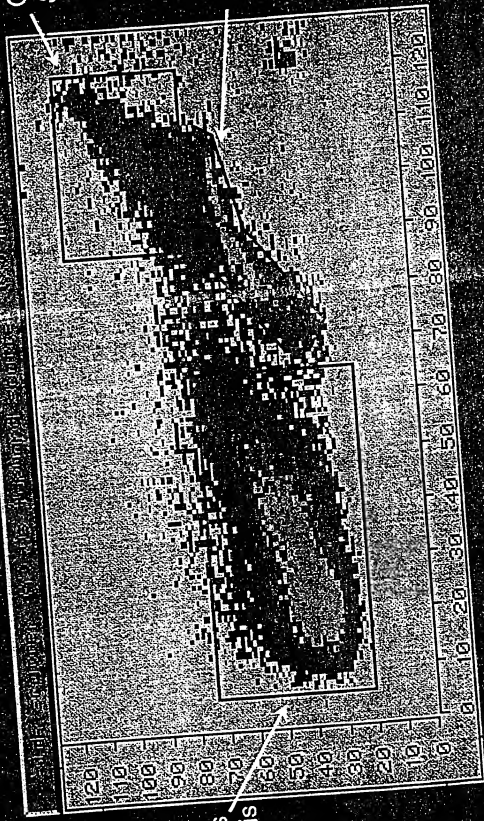
001180-870252560

FACS Enrichment for Gel Microdroplets Containing *S. div*



GMDs with
S. div

Empty
GMDs



Free cells
and debris

